Eliminating Vertically-transmitted HIV/AIDS while Improving Access to Treatment and Care for Women, Children and Adolescents in Jamaica

CDC Christie, RB Pierre

ABSTRACT

Background and Methods: To celebrate Jamaica's 50th birthday after receiving independence from Great Britain, we summarize our collaborative published research in the prevention, treatment and care of paediatric, perinatal and adolescent HIV/AIDS in Jamaica.

Results: Public access to antiretroviral therapy (ART) in Jamaica has shown that a "test and treat" strategy associated with "treatment for prevention" works for HIV-infected pregnant women by reducing their HIV-attributable morbidity and mortality and reducing mother-to-child transmission (MTCT) rates to < 2%, islandwide. These women experience significant psychosocial stress and targeted interventions are assisting them to improve their quality of life. HIV-exposed and infected children come from large families with high rates of teen pregnancies and significant financial challenges needing sustained interventions. HIV-exposed but uninfected Jamaican infants have higher rates of communityacquired infections, including lower respiratory tract infections, sepsis and gastroenteritis compared to community controls, although their growth rates are normal. In evaluation of replication capacity, viral control and clinical outcomes after vertical transmission in Jamaican mother-infant pairs, HLA-B57 was found to confer the advantage of restricted HIV replication primarily by driving and maintaining a fitness-attenuating mutation in p-24 Gag. Viral sequences from 52 MTCT Jamaican pairs were compared and 1475 sites of mother-infant amino acid divergence within Nef, Gag and Pol were identified, suggesting modest fitness cost with many CD8 mutations. HIV-infected Jamaican children are surviving into adolescence and adulthood, as a result of increased public access to ART and improved collaborative capacity in ART management. Successful transition of HIV-infected children through adolescence into adulthood requires a strong multidisciplinary team approach, including long-term ART management addressing non-adherence, drug resistance and toxicity, treatment failure and limited options for second line and salvage therapy, while attending to their sexual and reproductive health, psychosocial, educational and vocational issues and palliative care.

Conclusion: Over the past nine years, Jamaica has made excellent strides to eliminate vertically transmitted HIV/AIDS, while reducing the HIV-attributable morbidity and mortality in pregnant women and in HIV-infected children. Continued successful transition of HIV-infected children through adolescence into adulthood will require a strong multidisciplinary team approach.

Keywords: AIDS, children, HIV, Jamaica, pMTCT, Youth

Eliminando la Transmisión Vertical del VIH/SIDA Mejorando a la par el Acceso al Tratamiento y el Cuidado de las Mujeres, los Niños y los Adolescentes en Jamaica

CDC Christie, RB Pierre

RESUMEN

Antecedentes y Métodos: A fin de celebrar el 50 aniversario de Jamaica tras recibir la independencia de Gran Bretaña, resumimos nuestra investigación colaborativa publicada sobre la prevención, tratamiento y cuidado del VIH/SIDA pediátrico, perinatal y juvenil en Jamaica.

From: Department of Adolescent Medicine and Paediatrics, The University of the West Indies, Kingston 7, Jamaica.

Correspondence: Professor CDC Christie, Vaccines Infectious Diseases Centre, Department of Adolescent Medicine and Paediatrics, The University of the West Indies, Kingston 7, Jamaica. E-mail: Celia.ChristieSamuels@uwimona.edu.jm

Footnotes: Funded in part by the Global Fund for AIDS, Tuberculosis and Malaria, Clinton Health Access Initiative, Elizabeth Glaser Paediatric AIDS Foundation, Pfizer Foundation, United States National Institutes of Health, United States National Institutes of Child Health and Human Development, Embassy of Jamaica in Washington, DC, University of the West Indies and the Jamaican Ministry of Health.

397 Christie and Pierre

Resultados: El acceso público a la terapia antiretroviral (TAR) en Jamaica ha mostrado que una estrategia "test and treat" asociada con el "tratamiento para la prevención" funciona de manera efectiva con mujeres embarazadas infectadas por VIH, reduciendo la morbilidad y la mortalidad atribuibles al VIH, y disminuyendo las tasas de transmisión madre a niño (MTCT) a < 2% en toda la isla. Estas mujeres experimentan un estrés psicosocial considerable, y las intervenciones aplicadas están ayudándolas a mejorar su calidad de vida. Los niños expuestos e infectados por el VIH provienen de familias numerosas con altas tasas de embarazos adolescentes y considerables retos financieros. Se trata pues de familias que necesitan intervenciones sostenidas. Los infantes jamaicanos expuestos pero no infectados por el VIH tienen tasas más altas de infecciones adquiridas en la comunidad – incluyendo infecciones de las vías respiratorias bajas, sepsis y gastroenteritis – en comparación con los controles comunitarios, si bien sus tasas de crecimiento eran normales. Al evaluar la capacidad de replicación, el control viral, y los resultados clínicos tras la transmisión vertical en pares madre-infante jamaicanos, se halló que el HLA-B57 confería la ventaja de restringir la replicación del VIH mediante la conducción y mantenimiento de una mutación atenuante de la aptitud adaptativa (fitness) en p-24 gag. Las secuencias virales de 52 pares jamaicanos MTCT fueron comparadas, y se identificaron 1475 sitios de divergencia de aminoácido de madre-infante dentro de nef, gag y pol, lo cual sugiere un costo modesto de aptitud adaptativa con muchas mutaciones de CD8. Los niños jamaicanos infectados por VIH están sobreviviendo hasta llegar a ser adolescentes o adultos, como resultado del aumento del acceso público a la TAR, y al mejoramiento de capacidad colaborativa en el tratamiento de TAR. La transición exitosa de niños infectados con VIH a través de la adolescencia hasta la adultez requiere un enfoque multidisciplinarlo en equipo, incluyendo el tratamiento de TAR a largo plazo. Dicho tratamiento se dirige a la no adherencia, la resistencia a los medicamentos y la toxicidad, el fracaso del tratamiento y opciones limitadas para las terapias de segunda línea y de salvamento, a la par que se atiende a la salud reproductiva y sexual de los pacientes, a los problemas vocacionales, educacionales y psicosociales, y el cuidado paliativo.

Conclusión: En los últimos nueve años, Jamaica ha dado pasos extraordinarios para eliminar la transmisión vertical del VIH/SIDA, reduciendo la morbilidad y la mortalidad atribuibles al VIH en mujeres embarazadas y en niños infectados por VIH. La transición exitosa continuada de los niños infectados por VIH a través de la adolescencia hasta la adultez requerirá un fuerte enfoque multidisciplinarlo en equipo.

Palabras claves: SIDA, niños, VIH, Jamaica, PMTCT, juventud

West Indian Med J 2012; 61 (4): 397

BACKGROUND

Jamaica is a middle income developing island nation in the Caribbean, with a population of 2.8 million and an adult HIV seroprevalence of 1.7% in 2011, where an estimated 32 000 persons are currently living with HIV (1). The first case of AIDS was diagnosed in 1982 in Jamaica; since then reported cases have gradually increased so that the epidemic is now characterized as mixed, with features of a generalized epidemic in the population, but concentrated in those who are most at risk. The Jamaican HIV epidemic has had strong leadership from the National AIDS Programme over the years mainly through the efforts of Professor J Peter Figueroa with financial support from international partners (1-3). Children in Jamaica with HIV infection are now surviving into adolescence and young adulthood, as a result of increased public access to antiretroviral therapy (ART) and improved capacity in ART management.

Purpose

To celebrate Jamaica's 50th birthday after receiving political independence from the British, we summarize herein our

collaborative research in paediatric, perinatal and adolescent HIV/AIDS in Jamaica. We report our collaborative research efforts to virtually eliminate mother-to-child transmission of HIV/AIDS and the outcomes of interventions to improve treatment and care to children and youth living in Jamaica.

Leadership and Training

In 1986, the first case of paediatric HIV/AIDS was diagnosed by Celia Christie, then a Paediatric Infectious Diseases Fellow, visiting Jamaica from Yale University (4). Patricia Burke and Russell Pierre later established the Family Centre at the University Hospital of the West Indies (UHWI) where some children with HIV were being treated before ART became available. Reports of paediatric cases gradually increased islandwide and a hospital-based description by Tracy Evans-Gilbert *et al* reported HIV/AIDS as a leading cause of death in young children (5). In 2002, a team of government and academic healthcare personnel from The University of the West Indies (UWI) began collaborating together to address the Jamaican paediatric and perinatal epidemic through an International Leadership Award to

Christie (6-12). A five-point strategy was utilized comprising leadership and train-ing, preventing mother-to-child transmission, paediatric and maternal treatment and care, outcomes-based research activities, as well as collaboration locally, regionally and internationally (6-12).

The intervention began in the Greater Kingston Metropolitan Region with "The Kingston Paediatric and Perinatal HIV/AIDS - KPPAIDS Programme," and then extended islandwide to the "Jamaica Paediatric and Perinatal HIV/AIDS - JaPPAAIDS Programme". It was coordinated from each hub by nurse-managers working through the antenatal clinics, paediatric sites, obstetric hospitals, laboratories and the National AIDS Programme collaborating with the patient-families and the entire healthcare team (8). Training was ongoing to the multidisciplinary healthcare team and students through didactic lectures, conferences, small group discussions and also clinical preceptorships, including training of five clinical postdoctoral paediatric fellows and nine cohorts of graduating paediatricians from the UWI. A collaborative database was developed and a memorandum of understanding was implemented to guide our collaborations and the process for research publications. Data were collected prospectively, analysed, reported nationally and utilized to inform peer-reviewed publications of several outcomes-based and other research activities (2-69).

Preventing Mother-to-child Transmission of HIV and Perinatal HIV/AIDS

Harvey et al implemented the first efforts to prevent motherto-child transmission of HIV through the offering of nevirapine prophylaxis in a pilot study to HIV⁺ pregnant women and their babies, but reported limited follow-up and testing of the babies (13). In the first year of the KPPAIDS Programme, Johnson et al reported HIV testing of 53% of 7383 pregnant women to identify 107 HIV+ women in Greater Kingston, 75% of whom received zidovudine, or nevirapine; with repeat pregnancies and poor partner notification observed in over 30% (14). Whyte et al also noted that there was no significant difference in mother-to-child transmission rates between women presenting early (7.7%) and late presenters (10%, p = 0.897), although the overall MTCT rates were 8.2% (15). During the first three years, Christie et al reported that while modified short course zidovudine or nevirapine significantly decreased MTCT rates from 29% to 6%, highly active antiretroviral therapy (HAART) later decreased MTCT to < 2% in Greater Kingston and < 5% islandwide (7).

The Table shows the uptake of the pMTCT cascade from 2005 to 2011 in the Jamaican public sector, with > 95% of pregnant women now being tested for HIV, > 85% of women are receiving ARTs and 100% of babies are getting ART chemoprophylaxis and being offered full replacement formula feeds; the 2011 national pMTCT rates are reported as 1.19%, which is synonymous with the established targets

Table: Prevention of mother-to-child transmission of HIV in Jamaica

	2005	2006	2007	2008	2009	2010	2011
# ANC Attendees Tested	28,651 (96%)	28,446 (95%)	22,478 (95%)	29,119 (>95%)	30,076	26,697	27985
# HIV +ve pregnant women delivered	401	442	358	616	440	432	354
% of women getting ARVs	74%	84%	84%	84%	84%	86.3	85%
# of HIV – exposed infants	407	433	362	612	439	419	350
# Infants getting PMTCT	353 (87%)	403 (93%)	350 (97%)	605 (98%)	430 (98%)	408 (97%)	354 (101%)
Transmission Rate	10%	<10%	<5%	<5%	4.3%	4.6%	1.19%*

*Preliminary transmission rate 2011: 1.19%

Source - Ministry of Health, National AIDS Programme, 2012

for the virtual elimination of paediatric AIDS [$ie \le 2\%$] (16). The significant downward trend in recent years of reported new cases of paediatric HIV/AIDS and attributable deaths in

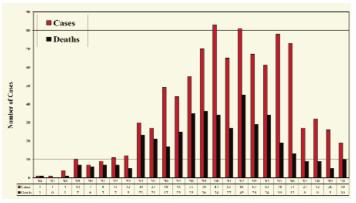


Figure: Paediatric AIDS cases and deaths in Jamaica, 1982–2010.

Source - Ministry of Health, National AIDS Programme, 2012

Jamaica is shown (Figure). Johnson *et al* reported further that the use of HAART when given to women in pregnancy along with the implementation of a comprehensive system of care including assessments of viral load and CD4 counts had greatly decreased HIV/AIDS-attributable maternal morbidity and mortality (17). Palmer *et al* noted that evaluation of the 15% noncompliance with ART in pregnancy revealed gaps in antenatal clinic attendance, prescribing and administering ART, hospital record documentation and follow-up of HIV exposed infants (18).

Psychosocial factors among HIV-infected and non-infected women in the peripartum period were evaluated by Weller *et al* who noted that significantly more HIV-infected women reported negative emotions, (*eg* guilt and tension), prayed, slept, or changed their eating habits, in addition to seeking the advice of a traditional faith healer (19).

Rodriquez *et al* noting a repeat pregnancy rate of 30-40% in over 800 HIV⁺ pregnant Jamaican women, reported the driving factors for repeat pregnancies included

399

issues relating to economics, fear of disclosure, rejection of contraceptives, reluctance to change lifestyle practices and the fact that having children was important in affirming the well-being and definition of womanhood, as a parent and as a partner (20). Rodriquez et al, further revealed in the same cohort that 70% of the women reported some kind of contraceptive use, with condoms accounting for 66%, although only half admitted consistent condom use and 16% had a tubal ligation. Their perceived inefficiencies of the healthcare system included delays in performing tubal ligations, inadequate access to contraceptive choices and advice, need for ongoing psychosocial support groups for stigma and discrimination (21). McCarthy also reported that psychotherapeutic interventions to mothers helped to identify, manage and treat their psychological and psychosocial issues and the resultant healing resulted in their accepting their HIV+ status, increased adherence to treatment, with gradual stabilization and self-efficacy to address the challenges of caring for themselves and their HIV-exposed infants, while enhancing the multidisciplinary approach to care (22). Moore et al reported that the challenges of a HIV voluntary counselling and testing (VCT) programme in the antenatal clinics included lack of quality control and lost opportunities for late presenters, while the successes included increased uptake of VCT and reduction of stigma (23). Hylton-Kong et al emphasized the role of the "contact investigator" as an essential member of the pMTCT team (24). Billings noted the increased prevalence of tattoos and body piercing in young women who are also sexually active and recognized the possible threat of transmission of blood-borne pathogens due to the unethical and unsterile practices of persons who are involved in this unregulated trade in Jamaica (25). Notwithstanding, Christie et al have reported that Jamaica along with several other Caribbean islands are well on their way to achieve their stated targets for the virtual elimination of vertical, or mother-to-child transmission of HIV/AIDS (26).

HIV-exposed Infants

Rodriquez et al in examining psychosocial dynamics and family structure noted that HIV-exposed and infected children come from large families, with a high prevalence of teen pregnancies and institutional care, creating a significant financial burden for Jamaica which needs sustained interventions to successfully address this (27). Steel-Duncan et al in following HIV-exposed infants early on in the Greater Kingston Metropolitan regional epidemic, reported that 97% received prophylaxis, 90% were not breastfed and 88% received cotrimoxazole prophylaxis, but challenges with stigma, non-compliance, breast milk substitution and follow-up care were noted and later addressed (28). Evans-Gilbert et al reported HIV-exposed infants were at risk of defaulting paediatric care if their mothers defaulted antenatal and post delivery care, therefore emphasizing the need for closer follow-up of these "at risk" mothers after delivery to identify

and define the HIV-status of their exposed infants (29). Mussi-Pinhata et al, in collaboration with the NISDI Perinatal Study Group, examined a cohort of HIV-exposed infants in Latin America and the Caribbean and reported the high risk of lower respiratory tract infections in HIV-exposed, uninfected infants (30). Fulford et al, in studying a similar cohort of HIV-exposed uninfected infants here in Jamaica, recognized 83% with at least one infection, comprising lower respiratory tract infections, sepsis, or acute gastroenteritis compared to normal HIV-negative and unexposed community controls, and although 30% of these HIV-exposed uninfected infants required hospitalizations, their growth trends were normal (31). Other collaborations with the NISDI Perinatal Study Group showed that Caesarean section before labour and membrane rupture was relatively safe for newborns of HIV-infected women (32). Similar NISDI collaborations reported the prevalence of congenital anomalies after the first trimester exposure to maternal ARTs was similar to that after the second and third trimester (33).

Immunology of Perinatal HIV in Mothers and Infants

International collaboration with Maggie Feeney et al to evaluate the immunology of perinatal HIV in Jamaican mother-infant pairs defined the HIV epitopes targeted by HIV-infected and non-infected Jamaican infants and reported qualitative differences in the CD8 response and deficient HIV-specific CD4 cells which may contribute to inability of young infants to limit HIV replication (34). In assessing the HLA B57-associated mutations on replication capacity, viral control and clinical outcomes after vertical transmission in Jamaican mother-infant pairs, it was concluded that HLA-B57 conferred its advantage of restricted HIV replication primarily by driving and maintaining a fitness-attenuating mutation in p-24 Gag (35). Frequency, epitope specificity and functional attributes of HIV-specific T-cells and sequence variation within B57-restricted epitopes were compared between Jamaican "spontaneous controllers" who maintained their normal CD4% and viral loads less than 3000 copies/ml without HAART and "treatment progressors" on HAART and it was concluded that among HLA B57 positive long-term survivors, spontaneous control of viraemia is not associated with a qualitatively or quantitatively superior Tcell response, but with uncompensated fitness-attenuating mutations in the viral capsid (36). Finally, viral sequences from 52 MTCT Jamaican pairs were compared and 1475 sites of mother-infant amino acid divergence within Nef, Gag and Pol were identified and it was concluded that there was modest fitness cost with many CD8 mutations (37).

Paediatric HIV/AIDS

Pierre *et al* characterized the natural history of paediatric HIV in a primarily ARV-naive population in Jamaica using the CDC criteria to define the disorders, reporting 88% of the children developing HIV from MTCT, 37% being severely symptomatic with *Pneumocystis jirovecii* pneumonitis and

tuberculosis, the most frequent opportunistic infections and Streptococcus pneumoniae, the most frequent bacterial pathogen (38-40). Using modified WHO guidelines, Pierre and Evans-Gilbert et al further reported that HAART was effective in decreasing opportunistic infections, hospitalizations and deaths in the HIV-infected children in both major cities in Greater Kingston and Montego Bay (41, 42-44). About 20–30% of the children were being placed on second line HAART (41, 43, 44). Adherence to HAART was 100% in children in institutional care compared to 78% among those in family based care, correlating with CD4% and according to White et al, non-adherence was related to older age of child, missing clinic appointments and nausea (45). Antiretrovirals and cotrimoxazole were reported to be safe in Jamaican children by Pryce et al, except for the few children early in the epidemic who changed ARV therapy due to anaemia from zidovudine, rash from nevirapine and indinavir associated haematuria and a few others who had drug toxicity with some biochemical abnormalities (46, 47). Roye et al found multiple drug genotypic resistance in the subset of children failing first-line ARVs (48). Heslop et al has reported a high genetic diversity of HIV in children with clade B, the predominant genotype in these children (49, 50).

At the height of the paediatric HIV epidemic, reports of tuberculosis (TB) increased, with HIV-infected children being statistically more likely to be older, have failure to thrive, digital clubbing, hepatosplenomegaly, adenopathy and negative Mantoux skin tests, to die, or have longer hospital stays when compared to the HIV-noninfected cohorts with TB (51). Geoghagen et al also reported concurrent outbreaks of infectious diseases, including tuberculosis, scabies and varicella in HIV-infected children living in a residential institution and emphasized the importance of education and appropriate immunization in children and the employees (52). Drug-resistant disseminated tuberculosis was also being seen (53). Immune reconstitution syndrome with BCG-lymphadenitis presumably from the Bacillus Calmette Guerin vaccine administered in the first week of life, was reported among several children after they had commenced HAART (54). Nicholson et al has reported disseminated histoplasmosis as a serious opportunistic infection in our cohort (55). Steel-Duncan et al noted renal manifestations including HIV-associated nephropathy and urinary tract infections linked to the common enteric pathogens (56). This work was extended by Byam et al who reported on the drug resistance patterns of these pathogens, recommending cotrimoxazole as a poor choice for empiric treatment of sepsis and urinary tract infections in this clinical setting (57).

Lewis *et al* reported anaemia, leucopenia and thrombocytopenia as not infrequent associations in ARV-naïve HIV-infected Jamaican children (58). Walker *et al* defined the prevalence of encephalopathy to be 23% in the Jamaican paediatric cohort associated with significant neurocognitive dysfunction (59). However, opportunistic enteric infections and infestations have not been reported in Jamaican children with HIV/AIDS who are on ARTs and cotrimoxazole (60). Partnerships with the private sector have enabled education of children in Jamaica's primary and secondary schools through debating competitions around moots of critical thinking, child rights, anti-discrimination, healthy lifestyles, parental and community responsibilities (61).

Adolescent HIV/AIDS

There are about 512 children and adolescents living with HIV/AIDS who are currently enrolled in treatment and care in Jamaica, most having acquired HIV from MTCT; 88% (451) are currently receiving HAART using modified WHO guidelines and of these, 73% (332) are < 12 years old and 27% (119) are aged > 13 years. Among the 451 on HAART, 70% (327) are on first line HAART, 29% (121) are on second line HAART and 0.7% (3) are receiving salvage therapy. These children and youth are mostly healthy and ambulatory. As a result of the strong collaborative partnerships and interventions by the Ministry of Health and the UWI, the significant downward trend in recent years of reported new cases of HIV/AIDS and deaths in children in Jamaica is shown (Figure).

Walker, and Harrison et al, have reported that adolescents with HIV are increasingly being recognized from the now ageing population of children who acquired HIV by vertical transmission and those who acquired HIV horizontally, from consensual, or forced sexual intercourse (62, Moore et al recently reported that among 115 adolescents aged 12-24 years being treated for HIV/AIDS at the UHWI, 98% received formal education, 46% of those > 18 years were employed, 71% acquired HIV by MTCT and 22% sexually, several had sexually transmitted infections and several became pregnant; it was concluded that this heterogeneous cohort of HIV-infected maturing youth had sexual/reproductive health, psychosocial, educational and vocational challenges that required a multidisciplinary team approach to address (64). Lewis-O'Connor et al have noted bone metabolic disease as a recognized cause of pathological fractures in HIV-infected adolescents on HAART; it is clear that screening with DEXA scan is desirable but financially constrained in resource-limited settings and so the management for maturing perinatally infected children should include supportive measures to include optimized ART regimes, calcium and vitamin D supplementation, improved diet and exercise (65). Dunkley-Thompson also emphasized the need for healthcare providers to recognize and report the long-term non-progressors among the missed population of perinatally acquired HIV-infected teens to link them to appropriate treatment, care and prevention programmes (66). Pilgrim et al have noted that the majority of HIV-infected adolescents lived with parents or guardians, suggesting support despite stigma and discrimination (67). Moore also reported that among the adolescents with horizontal acquisition of HIV/AIDS, about half had acquired HIV infection forcibly (64). In addition to the legal interventions, post-exposure prophylaxis and psychotherapy must be included in the treatment for youth who are the victims of sexual assault (68–69).

Adolescents and youth in Jamaica continue to be at risk of HIV infection due to early sexual debut, 13 years in males and 15 years in females (1). Despite the increased perception of HIV risk in this population, there is no significant change in youths reporting multiple sex partnerships or the age of first sex (1). Contributing factors include poor condom negotiating skills by females, early sexual initiation with older men, high prevalence of sexual abuse of adolescent females and increased detection through voluntary counselling and testing for all antenatal clinic attendees (1). Youth who are being infected from sexual transmission are also at risk for possible onward transmission of HIV to their partners. There is also the growing population of youth who themselves were saved from MTCT who are now growing up in the same environment that contributed to their parents' infection and are now at risk for acquiring HIV from sexual transmission.

Operational Challenges

Although Jamaica has reached the virtual elimination target in that the rate for HIV⁺ infants born to HIV-infected mothers is now 2% or less, an incidence of MTCT of 0.3 cases per 1000 live births also has to be attained and both targets be maintained for three years, for the country to be officially certified. There are several concrete operational and other challenges that still need to be overcome to achieve these goals. These include closing the gap to improve ART uptake in the 15% of women who still elude the pMTCT programme by presenting late, or not at all, or who are not HIV tested, or do not reveal their HIV+ serostatus on the labour ward. HIV rapid testing, although available, needs to be routinely performed on the labour ward for women who do not have a HIV test result in their hospital chart. Partners need to be brought in for counselling and HIV testing. The repeat pregnancy rates of about 25-30% each year and related risky sexual behaviours need to be addressed. The 15% of HIV-exposed infants who are "lost to follow-up" need to be identified for definitive HIV polymerase chain reaction (PCR) testing; although we know they are probably HIVnegative because their mothers did participate in the pMTCT cascade and the infants benefitted from ART prophylaxis and replacement formula, further they would have presented to the health sector with AIDS-defining illnesses, if they were HIV-infected. Their linkage to and retention in care, needs a better integrated and decentralized service delivery to expand access to HIV diagnosis and treatment interventions and thereby reduce loss to follow-up. HIV testing data for the women from the private sector of about 30% need to be accounted for by improving public private partnerships, although it is known that these women are eventually accounted for among the HIV+ deliveries, as they deliver their babies in the public hospitals. Stigma and discrimination in the health sector and community still need to be proactively addressed.

The main operational challenges of "treatment and prevention" for HIV-infected children and adolescents in Jamaica are many. These may include the age appropriate issues of providing "disclosure" of HIV status to and from the child or adolescent for their own health and the health of the youth's partners. The ethics of HIV testing in children and youth, eg confidentiality, informed age appropriate consent, or assent, inappropriate disclosure, eg in schools need to be considered. Adherence counselling and psychosocial support needs to be made available for appropriate treatment, prevention and care. Paediatric drug optimization must be considered along with appropriate management of drug toxicity as it relates to first line and second line ART and salvage therapy. Sexual and reproductive health support needs to be provided as this relates to sex, sexual abuse, pregnancy and sexually transmitted diseases. Continued successful transition of HIV-infected children through adolescence into adulthood will require a strong multidisciplinary team approach, including long-term ART management addressing nonadherence, drug resistance and toxicity, treatment failure and limited options for second line and salvage therapy, while attending to their sexual and reproductive health, psychosocial, educational and vocational issues and palliative care. The tattooing and body piercing industry in Jamaica needs to be regulated to ensure proper infection control and prevent exposure to blood borne pathogens, including HIV. grammes need to be put in place for at risk youth to reduce their risky behaviours, eg street children, drug abusers. Consideration needs to be made for integration of services for maximal impact, eg maternal, neonatal, child, sexual health, men's health and family planning. These considerations need to be addressed amidst the global economic climate of attrition of funding (eg Global Fund, CHAI, UNITAID).

CONCLUSION

Over the past nine years, Jamaica has made excellent strides in the public sector to eliminate vertically transmitted HIV/AIDS, while reducing the HIV-attributable morbidity and mortality in pregnant women and in HIV-infected children.

Public access to ART in Jamaica has shown that a "test and treat" strategy associated with "treatment for prevention" works for HIV-infected pregnant women by reducing their HIV-attributable morbidity and mortality and reducing mother-to-child transmission rates to < 2% islandwide. These women experience significant psychosocial stress and targeted interventions are assisting them to improve their quality of life. HIV-exposed and infected children come from large families with high rates of teen pregnancies and significant financial challenges needing sustained interventions. HIV-exposed but uninfected Jamaican infants have higher rates of community-acquired infections, including lower res-

piratory tract infections, sepsis and gastroenteritis compared to community controls, although their growth rates are normal. In evaluation of replication capacity, viral control and clinical outcomes after vertical transmission in Jamaican mother-infant pairs, HLA-B57 was found to confer its advantage by restricting HIV replication primarily by driving and maintaining a fitness-attenuating mutation in p-24 Gag. Viral sequences from 52 MTCT Jamaican pairs were compared and 1475 sites of mother-infant amino acid divergence within Nef, Gag and Pol were identified, suggesting modest fitness cost with many CD8 mutations. HIV-infected Jamaican children are surviving into adolescence and adulthood, as a result of increased public access to ART and improved collaborative capacity in ART management. Successful transition of HIV-infected children through adolescence into adulthood requires a strong multidisciplinary team approach, including long-term ART management addressing non-adherence, drug resistance and toxicity, treatment failure and limited options for second line and salvage therapy, while attending to their sexual and reproductive health, psychosocial, educational and vocational issues and palliative care.

REFERENCES

- Jamaica HIV/AIDS/STI National Strategic Plan 2012–2017. Kingston: Ministry of Health.
- Figueroa JP. The HIV epidemic in the Caribbean: Meeting the challenges of achieving universal access to prevention, treatment and care. West Indian Med J 2008; 57: 195–203.
- Figueroa JP. An overview of HIV/AIDS in Jamaica. Strengthening the response. West Indian Med J 2004; 53: 277–82.
- Christie CD, Bain B, Pierre R, Smikle M, Evans-Gilbert T, Fredericks J et al. HIV/AIDS in women, infants, children and adolescents in Jamaica: a further call to action. West Indian Med J 2001; 50: 258–62.
- Evans-Gilbert T, Hambleton I, McKenzie CA, Samms-Vaughan M. Paediatric HIV/AIDS in Jamaica – a hospital-based study. West Indian Med J 2002; 51: 74–9.
- Christie CDC. A paediatric and perinatal HIV/AIDS leadership initiative in Kingston, Jamaica. West Indian Med J 2004; 53: 283–92.
- Christie CDC, Steel-Duncan J, Palmer P, Pierre R, Harvey K, Johnson N et al. Paediatric and perinatal HIV/AIDS in Jamaica: An international leadership initiative, 2002–2007. West Indian Med J 2008: 57: 204–15.
- Palmer P, Anderson-Allen MM, Billings CC, Moore JT, McDonald-Kerr C, Steel-Duncan JC et al. Nursing interventions in the Kingston Paediatric and Perinatal HIV/AIDS Programme. West Indian Med J 2004: 53: 327–31.
- Safrit JT, Wilfert CM. Paediatric HIV/AIDS in Jamaica: Present success and future challenges. West Indian Med J 2004; 53: 271–3.
- Safrit J, Wilfert C, eds. Paediatric and perinatal HIV/AIDS in Jamaica [Special issue]. West Indian Med J 2004; 53: 271–365.
- Vermund SH, Krogstad PA, guest eds. Prevention of mother-to-child HIV transmission and management of paediatric HIV infection in Jamaica. West Indian Med J 2008; 57: 187–8.
- Vermund SH, Krogstad PA, guest eds. Paediatric and perinatal HIV/ AIDS in Jamaica [Special issue]. West Indian Med J 2008; 57: 187–320.
- Harvey KM, Figueroa JP, Tomlinson J, Gebre Y, Forbes S, Toyloy T et al. An assessment of mother-to-child transmission prevention in 16 pilot antenatal clinics in Jamaica. West Indian Med J 2004; 53: 293-6.
- 14. Johnson N, Mullings A, Harvey K, Alexander G, McDonald D, Smikle M et al. HIV Seroprevalence, uptake of interventions to reduce motherto-child transmission and birth outcomes in greater Kingston, Jamaica.

- West Indian Med J 2004: 53: 297-302.
- 15. Whyte K, Pierre RB, Anderson-Allen M, Williams E, Harvey K, Palmer P et al. The pregnancy outcomes of HIV positive mothers who present late to high-risk-antenatal clinics in Jamaica. Electronic poster presentation at: IAS 2009; 5th IAS Conference on HIV Pathogenesis Treatment and Prevention; July 19–22, 2009; Cape Town, South Africa. Abstract A-155-0107-02739, electronic poster CDC022.
- Christie CDC, Palmer P, Lewis K, Pierre R, Moore J, Anderson-Allen M et al. Jamaica is achieving MDG elimination targets for vertically transmitted HIV/AIDS. Abstract and poster presented at: 6th IAS Conference on HIV Pathogenesis, Treatment and Prevention; July 17–20, 2011; Rome, Italy. Abstract: TUPE 290; 2763; A-361-0139-02763).
- 17. Johnson N, Palmer P, Samuels LA, Morgan O, Onyonor A, Anderson M et al. Evolving care of HIV-infected pregnant women from nevirapine to HAART. West Indian Med J 2008; 57: 216–22.
- 18. Palmer P, Moore J, Anderson-Allen M, Gilman S, Billings C, Heron-Moore T, et al. Gap analysis of ART uptake in HIV-infected pregnant women towards achieving the elimination goal in Jamaica. Abstract and poster WEPE 099 presented at: X1X International AIDS Conference; July 22–27, 2012, Washington DC.
- Weller PD, Hambleton I, Chambers C, Bain S, Christie CDC, Bain B. The voices of the women: Feedback from women of child-bearing age who are living with HIV can help improve efficacy of psycho-social interventions. West Indian Med J 2008; 57: 274–81.
- Rodriquez LB, White S, Frankson M, Billings C, Christie CDC. Factors influencing repeat pregnancies in HIV-positive mothers in Jamaica. Abstract and poster TU 0832 presented at: XVII International AIDS Conference; August 3–8, 2008, Mexico City.
- Rodriquez LB, White S, Frankson M, Billings C, Christie CDC. Issues
 of family planning among HIV-positive women in Jamaica. Abstract
 and electronic poster CDC 168 presented at: IAS 2009; 5th IAS
 Conference on HIV Pathogenesis Treatment and Prevention; July
 19–22, 2009; Cape Town, South Africa.
- McCarthy R. Psychological and psychosocial issues related to pMTCT management among referred patients. Abstract and poster MOPE 433 presented at: X1X International AIDS Conference; July 22–27, 2012. Washington DC.
- Moore J, Palmer P, Anderson-Allen M, Billings C, McDonald-Kerr C, the Kingston Paediatric and Perinatal HIV/AIDS Study Group. Voluntary counselling and testing in antenatal clinics in Greater Kingston, Jamaica. West Indian Med J 2008; 57: 269–73.
- Hylton-Kong T, Alveranga T, Norrine T, Harris H, Daubon G. Contact investigation in the prevention of mother-to-child transmission of HIV, comparing urban and rural outcomes in Jamaica. West Indian Med J 2008; 57: 282–6.
- Billings C, the Kingston Paediatric and Perinatal HIV/AIDS Study Group. Tattooing and perinatal HIV/AIDS in Jamaica. West Indian Med J 2008; 57: 312–4.
- 26. Christie CDC, Caffe S, Wilson V, Del Riego A, Jack N, Bain B et al. Protecting our future generations: Born free of HIV in the Caribbean. Abstract A-361-0139-02644 presented at: 6th International Conference on HIV Pathogenesis, Treatment and Prevention; July 17–20, 2011; Rome, Italy.
- Rodriguez B, Steel-Duncan JC, Pierre R, Evans-Gilbert T, Hambleton I, Palmer P et al. Socio-demographic factors of HIV-exposed and HIVinfected Jamaican children. West Indian Med J 2004; 53: 303–07.
- Steel-Duncan J, Pierre R, Evans-Gilbert T, Rodriquez B, Smikle M, Palmer P, et al. Uptake of interventions, outcomes and challenges in caring for HIV-exposed infants in Kingston, Jamaica. West Indian Med J 2004; 53: 308–14.
- Evans-Gilbert T, Reid G, Spence K, Christie CDC. Adherence in children begins in pregnancy: A nested case control study of HIV-exposed infants born in Western Jamaica from 2008–2010. Abstract and poster # MOPE 057 presented at: X1X International AIDS Conference; July 22–27, 2012; Washington DC.
- Mussi-Pinhata MM, Motta F, Freimanis Hance L, de Souza R, Szyld E, Succi RCM et al. Lower respiratory tract infections among HIV-

- exposed, uninfected infants. Int J Infec Dis 2010; **14 (Suppl 3):** e176–82. Epub 2010 May 8.
- Fulford TA, Palmer P, Pierre R, Christie CDC. Infectious disease morbidity and growth patterns of HIV-exposed uninfected infants in Jamaica: an epidemiological cohort study. Abstract and poster presented at: X1X International AIDS Conference; July 22–27, 2012; Washington DC.
- Kreitchmann R, Cohen RA, Stoszek SK, Pinto JA, Losso M, Pierre R et al. Mode of delivery and neonatal respiratory morbidity among HIVexposed newborns in Latin America and the Caribbean: NISDI Perinatal-LILAC Studies. Int J Gynaecol Obstet 2011; 114: 91–6. Epub 2011 May 26. DOI: 10.1016/j.ijgo.2011.02.008.
- Joao EC, Calvet GA, Krauss MR, Freimanis HL, Ortiz J, Ivalo SA et al. Maternal antiretroviral use during pregnancy and infant congenital anomalies: the NISDI perinatal study. J Acquir Immune Defic Syndr 2010; 53: 176–85.
- Huang S, Dunkley-Thompson J, Tang Y, Macklin EA, Steel-Duncan J, Singh-Minott I et al. Deficiency of HIV-Gag-specific T cells in early childhood correlates with poor viral containment. J Immunol 2008; 181: 8103-11.
- Schneidewind A, Tang Y, Brockman MA, Ryland EG, Dunkley-Thompson J, Steel-Duncan JC et al. Maternal transmission of HIV escape mutations subverts HLA-B57 immuno-dominance but facilitates viral control in the haplo-identical infant. J Virol 2009; 83: 8616–27. Epub 2009 Jun 10.
- Tang Y, Huang S, Dunkley-Thompson J, Steel-Duncan J, Ryland EG, St John MA et al. Correlates of spontaneous viral control among longterm survivors of perinatal HIV-1 infection expressing human leucocyte antigen – B57. AIDS 2010; 24: 1425–35.
- Ryland EG, Tang Y, Christie CD, Feeney ME. Sequence evolution of HIV-1 following mother-to-child transmission. J Virol 2010; 8: 12437–44. Epub 2010 Sep 22.
- Pierre R, Steel-Duncan J, Evans-Gilbert T, Rodriguez B, Palmer P, Smikle M et al. CDC-defined diseases and opportunistic infections in Jamaican children with HIV/AIDS. West Indian Med J 2004; 53: 315-21
- Pierre R, Ramsay DH, Loudon M. HIV/AIDS and affected Jamaican children – a vulnerable generation. Caribbean childhoods: from research to action: Journal of the Children's Issues Coalition, 2005; 2: 82–99.
- Pierre R, Bailey KR, Dicks BA, Ramsay DH. HIV/AIDS and the Jamaican child – Implications for research and action. Caribbean Journal of Social Work 2005; 4: 59–72.
- Pierre R, Steel-Duncan J, Evans-Gilbert T, Rodriguez B, Moore J, Palmer P et al. Effectiveness of interventions in treating paediatric HIV/AIDS in Jamaican children. West Indian Med J 2008; 57: 223-30.
- Evans-Gilbert T, Steel-Duncan J, Pierre R, Rodriguez B, Palmer P, Figueroa JP et al. HIV-related mortality in Jamaican children. West Indian Med J 2008; 57: 265–8.
- Evans-Gilbert T, Pierre R, Steel-Duncan J, Rodriguez B, Whorms S, Palmer P et al. Anti-retroviral drug therapy in HIV-infected Jamaican children. West Indian Med J 2004; 53: 322-6.
- 44. Pierre R, Palmer P, Moore J, Christie CDC. Long-term outcomes of antiretroviral therapy (ART) in Jamaican children: an epidemiological cohort study. Abstract and poster #MOPE 210 presented at: X1X International AIDS Conference; July 22–27, 2012; Washington DC.
- White YRG, Pierre RB, Steel-Duncan J, Palmer P, Evans-Gilbert T, Moore J et al. Predictors of adherence to antiretroviral therapy in Jamaican children with HIV/AIDS. West Indian Med J 2008; 57: 231–7.
- Pryce C, Pierre R, Steel-Duncan J, Evans-Gilbert T, Palmer P, Moore J et al. Safety of antiretroviral drug therapy in Jamaican children with HIV/AIDS. West Indian Med J 2008; 57: 238–45.
- 47. Steel-Duncan J, Pierre R, Gabay L, Christie CDC. Nevirapine-associated rash in a Jamaican child. West Indian Med J 2004; 53: 356–8.
- Roye ME, Ramkisoon A, Amarakoon II, Hamilton CC, Eyzaguirre LM, Pierre RP et al. Multiple drug resistance in HIV-1 infected Jamaican children. West Indian Med J 2010; 59 (Suppl 4): 24–5.

- Heslop O, Smikle MF, Deer D, Christian N, Vickers IE, Harvey KM et al. Human immunodeficiency virus type-1 (HIV-1) subtypes in Jamaica. West Indian Med J 2005; 54: 279–82.
- Heslop OD, Smikle MF, Vickers IE, Christian NA, Harvey KM, Figueroa JP et al. High genetic diversity of HIV-1 in Jamaica. West Indian Med J 2009; 58: 195–200.
- Geoghagen M, Farr JA, Hambleton I, Pierre R, Christie CDC. HIV and TB co-infections in Jamaican children. West Indian Med J 2004; 53: 339–45.
- Geoghagen M, Pierre R, Evans-Gilbert T, Rodriguez B, Christie CDC. Tuberculosis, scabies and chicken pox outbreaks in an orphanage for children with HIV/AIDS in Jamaica. West Indian Med J 2004; 53: 346-51.
- Singh-Minott I, Pierre R, Olugbuyi O, Dunkley-Thompson J, Haughton D, Christie CDC. Isoniazid-resistant disseminated tuberculosis in a Jamaican infant with HIV/AIDS. West Indian Med J 2008; 57: 298–301
- Dunkley-Thompson J, Pierre R, Steel-Duncan J, Palmer P, Davis D, Figueroa P et al. Bacille Calmette-Guérin lymphadenitis following antiretroviral therapy in Jamaican infants with rapid progressor HIV disease. West Indian Med J 2008; 57: 302–06.
- Nicholson AM, Rainford L, Elliott V, Christie CDC. Disseminated histoplasmosis and AIDS at the University Hospital of the West Indies. West Indian Med J 2004; 53: 126–30.
- Steel-Duncan J, Miller M, Pierre R, Dunkley-Thompson J, Palmer P, Evans-Gilbert T et al. Renal manifestations of HIV/AIDS in Jamaican children. West Indian Med J 2008; 57: 246–52.
- 57. Byam PR, Pierre RB, Christie CDC, Andiman WA, Pettigrew M, the Kingston Paediatric and Perinatal (KPAIDS) Study Group. Antibiotic resistance among pathogens causing disease in Jamaican children with HIV/AIDS. West Indian Med J 2010; 59: 386–92.
- 58. Lewis K, Pierre R, Frankson A, Palmer P, Christie C, KPPAIDS Research Group. Haematological manifestations of human immunodeficiency virus (HIV) in an antiretroviral-naïve paediatric cohort. Abstract CDB0016 presented at: AIDS 2008. Proceedings of the XVII International AIDS Conference; August 3–8, 2008; Mexico City, Mexico.
- Walker S, Pierre R, Christie CDC, Chang-Lopez S, Kingston Paediatric and Perinatal HIV/AIDS Programme (KPAIDS) Research Group. Neuro-cognitive function in HIV-positive children in a developing country. Oral platform presentation at: Paediatric HIV and ART Complications. XVIII International AIDS Conferenc; July 18–23, 2010; Vienna, Austria, Abstract # 11032.
- Barrett D, Steel-Duncan J, Eldemire-Shearer D, Christie CDC, Lindo J. Absence of opportunistic parasitic infestations among children living with HIV/AIDS in children's homes in Jamaica. West Indian Med J 2008; 57: 253–6.
- Brissett D, Griffiths-Irving J. Speak up! Speak out! Building HIV and AIDS awareness among Jamaican children. West Indian Med J 2008; 57: 315–20.
- Walker E, Mayes B, Ramsay H, Hewitt H, Bain B, Christie CDC. Socio-demographic and clinical characteristics of Jamaican adolescents with HIV/AIDS. West Indian Med J 2004; 53: 332–8.
- Harrison A, Pierre R, Moore J, Davis D, Christie CDC. Clinical manifestations of adolescents with HIV/AIDS in Jamaica. West Indian Med J 2008; 57: 257–64.
- 64. Moore JA, Palmer P, Pierre R, Christie CDC. An emerging generation: socio-demography and sexual health in HIV positive youth in Jamaica to guide therapy. Abstract and poster WEPE434 presented at: X1X International AIDS Conference; July 22–27, 2012; Washington DC.
- Lewis-O'Connor K, Pierre RB, Moore JT, Christie CDC. Pathological fractures in adolescents on highly active antiretroviral therapy. Abstract and poster presented at: Caribbean HIV/AIDS Conference; November 2011; Nassau, Bahamas.
- Dunkley-Thompson J, Figueroa JP, Christie CDC. The "missed" population of perinatally-acquired HIV-infected adolescents in Jamaica. West Indian Med J 2006; 55: 295–7.

- Pilgrim N, Kershaw T, Pierre RB, Moore J, Palmer P, Davis D et al. Predictors of HIV/AIDS confirmation and differences by guardian status in HIV⁺ adolescents in Jamaica. West Indian Med J 2008; 57: 287–92.
- 68. Steel-Duncan JC, Pierre R, Evans-Gilbert T, Rodriguez B, Christie CDC. HIV/AIDS following sexual assault in Jamaican children and
- adolescents a case for post exposure prophylaxis for HIV. West Indian Med J 2004; 53: 352-5.
- 69. Lowe GA, Gibson RC, Christie CDC. HIV infection, sexual abuse and social support in Jamaican adolescents referred to a psychiatric service. West Indian Med J 2008; **57:** 307–11.

The Fertility Management Unit, Department of Obstetrics and Gynaecology and Child Health, The University of the West Indies, Kingston, Jamaica.

- Intra-uterine Insemination (IUI)
- In Vitro Fertilization and Embryo Transfer
- Cryopreservation and Frozen Embryo Transfer
- Semen Analysis
- Testicular Sperm Aspiration (TESA)

Telephone: (876) 970-2388

Fax #: (876) 927-0100

E-mail: joseph.frederick@uwimona.edu.jm